METHOD OF PREVENTING DEPLETION OF NON-AUTOLOGOUS HEMATOPOIETIC

CELLS AND ANIMAL MODEL SYSTEM FOR USE THEREOF U.S. Serial No. 08/704,445

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AMENDED CLAIMS

- 1. A method of preventing depletion in an animal of non-autologous hematopoietic cells comprising decreasing the number of endogenous macrophages to a level effective to substantially prevent depletion of the non-autologous hematopoietic cells.
- 2. The method according to claim 1 wherein the non-autologous hematopoietic cells are injected into the animal.
- 3. The method according to claim 1 wherein the cells are made by hematopoietic tissue engrafted into the animal.
- 4. The method according to claim 1 wherein the macrophages are decreased by administering to the animal an effective amount of an agent which decreases the level of endogenous macrophages.
- 5. The method according to claim 4 wherein the agent is liposome-encapsulated dichloromethylene diphosphonate.
- 6. The method according to claim 1 wherein the macrophages are decreased genetically.
 - 7. The method according to claim 1 wherein the animal is immunocompromised.
- 8. The method according to claim 7 wherein the animal is immunocompromised due to infection with an immunodeficiency virus.

- 9. The method according to claim 8 wherein the animal is human and the virus is human immunodeficiency virus.
- 10. The method according to claim 7 wherein the animal is immunocompromised due to radiation therapy.
- 11. The method according to claim 7 wherein the animal is immunocompromised due to chemotherapy.
- 12. The method according to claim 7 wherein the animal is selected from the group consisting humans, mice, scid/scid mice, SCID-hu mice, and CID horses.
- 13. The method according to claim 12 wherein the animal is a SCID-hu Thy/Liv mouse.
- 14. The method according to claim 7 wherein the animal is transplanted with non-autologous hematopoietic tissue.
- 15. The method according to claim 7 wherein the non-autologous hematopoietic cells are injected into the animal.
- 16. The method according to claim 7 wherein the agent is liposome-encapsulated dichloromethylene diphosphonate.
- 17. The method according to claim 15 wherein the animal is a human and the non-autologous hematopoietic cells are injected.
- 18. A method of treating an immunocompromised animal comprising administering to the animal an effective amount of non-autologous hematopoietic cells and decreasing endogenous macrophages to a level sufficient to prevent substantial depletion of the non-autologous hematopoietic cells.

- 19. A non-human mammal comprising human hematopoietic cells wherein the mammal contains a decreased level of endogenous macrophages sufficient to prevent substantial depletion of non-autologous hematopoietic cells.
- 20. The non-human mammal according to claim 19 wherein the mammal is immunocompromised.
- 21. The non-human mammal according to claim 19 wherein the mammal contains engrafted human hematopoietic tissue.
- 22. The non-human mammal according to claim 19 wherein the non-autologous hematopoietic cells are produced by the engrafted tissue.
- 23. The mammal according to claim 19 wherein the mammal is selected from the group consisting of SCID/SCID mice, SCID-hu Thy/Liv mice and CID horses.
- 24. A method of restoring hematopoietic cells to an immunocompromised human comprising the steps of administering an effective amount of human peripheral blood cells in conjunction with decreasing endogenous macrophages.
- 25. The method according to claim 24 wherein the immunocompromised human is infected with human immunodeficiency virus.
- 26. The method according to claim 25 wherein the peripheral blood cells are hematolymphoid.
 - 27. The method according to claim 26 wherein the blood cells are T cells.
 - 28. The method according to claim 26 wherein the blood cells are CD4⁺ T cells.

- 29. The method according to claim 25 wherein the peripheral blood cells are administered by direct injection into the blood stream of the human.
- 30. The method according to claim 25 wherein the peripheral blood cells are administered by bone marrow transplantation of hematopoietic stem cells into the human.
- 31. A method of improving engraftment efficiency for transplantation of a population of non-autologous hematopoietic stem cells in a host animal having an endogenous hematopoietic stem cell population, comprising the steps of ablating the endogenous hematopoietic stem cell population of the host animal and transplanting the non-autologous hematopoietic stem cells into the host animal in conjunction with decreasing endogenous macrophages in the host animal.